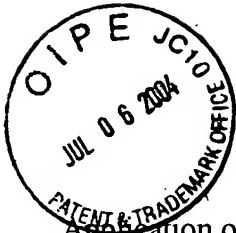


07-07-04

3412/1A/US (PHA 4140.6)
PATENT

1654
JPW



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of Mark G. Obukowicz et al.

Art Unit 1654

Serial No. 10/022,320

Filed December 13, 2001

Confirmation No. 4956

For SELECTIVE COX-2 INHIBITION FROM NON-EDIBLE PLANT EXTRACTS

Examiner Randall O. Winston

July 6, 2004

TO THE COMMISSIONER OF PATENTS AND TRADEMARKS,

SIR:

**LETTER TO THE PATENT AND TRADEMARK OFFICE REGARDING
INFORMATION DISCLOSURE STATEMENT**

On April 17, 2002, applicants submitted an Information Disclosure Statement listing 96 references. Applicants, however, did not have a copy of one of these references (reference 94) available to submit with the Information Disclosure Statement at that time. Applicants have now obtained and have enclosed a copy of the missing reference and a copy of the Information Disclosure Statement already submitted with this letter.

Respectfully submitted,

Timothy B. McBride, Reg. No. 47,781
SENNIGER, POWERS, LEAVITT & ROEDEL
One Metropolitan Square, 16th Floor
St. Louis, Missouri 63102
(314) 231-5400

TBM/sxm

*Enclosures

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PTO/SB/08A INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)				Complete if Known	
				Application Number	10/022,320
				Filing Date	12-13-2001
				Confirmation Number	4956
				First Named Inventor	M. Obukowicz
				Group Art Unit	1614
Sheet	1	of	9	Attorney Docket No.	3412/1A/US (PHA 4140.6)

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	U.S. Patent Document		Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY
		Number	Kind Code ² (if known)		
	1	4,613,591	B1	Aburada et al.	09-23-1986
	2	5,380,738	B1	Norman et al.	01-10-1995
	3	5,344,991	B1	Reitz et al.	09-06-1994
	4	5,393,790	B1	Reitz et al.	02-28-1995
	5	5,466,823	B1	Talley et al.	11-14-1995
	6	5,521,207	B1	Graneto	05-28-1996
	7	5,633,272	B1	Talley et al.	05-27-1997
	8	5,753,688	B1	Talley et al.	05-19-1998
	9	5,760,068	B1	Talley et al.	06-02-1998
	10	5,811,425	B1	Woods et al.	09-22-1998
	11	5,932,598	B1	Talley et al.	08-03-1999

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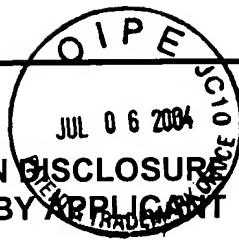
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Sheet

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FOREIGN PATENT DOCUMENTS

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		Office	Number ⁴	Kind Code ² (if known)			
	12	WO	94/15932	A1	G.D. Searle & Co.	07-21-1994	
	13	WO	95/15316	A1	G.D. Searle & Co.	06-08-1995	
	14	WO	00/74696	A1	Oxford Natural Products PLC	12-14-2000	
	15	JP	4208222		Tsumura & co.	07-29-1992	X
	16	JP	4005237	A	Nonogawa Shoji:kk (abstract)	01-09-1992	
	17	JP	2000236835		Seika et al. (abstract)	09-05-2000	
	18	EP	248215	B1	Popp et al.	12-09-1987	
	19	WO	94/12165	A	Wellcome et al.	06-09-1994	
	20	WO	94/14780	A	Wellcome et al.	07-07-1994	
	21	WO	93/13055	A	Glaxo Wellcome Inc.	07-08-1993	
	22	WO	95/11231	A	Bergmanis et al.	04-27-1995	

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OTHER ART - NON PATENT LITERATURE DOCUMENTS				
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.) date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ⁶	
	23	KASE, et al., Mechanisms by which Hange-shashin-to reduces prostaglandin E2 levels, <i>Biol. Pharm. Bull.</i> , 1998, pp. 1277-1281, Vol. 21, No. 12.		
	24	RINGBOM et al., Ursolic Acid from <i>Plantago major</i> , a Selective Inhibitor of Cyclooxygenase-2 Catalyzed Prostaglandin Biosynthesis, <i>J. Nat. Prod.</i> , 1998, pp. 1212-1215, Vol. 61, No. 10.		
	25	PENNISI, E., Building a Better Aspirin, <i>Science</i> , 1998, pp. 1191-1192, Vol. 280.		
	26	RESCH et al., 5-Lipoxygenase and Cyclooxygenase-1 Inhibitory Active Compounds from <i>Atractylodes Lancea</i> , <i>J. Nat. Prod.</i> , 1998, pp. 347-350, Vol. 61.		
	27	RUBIN, et al., Pharmacokinetics, Safety, and Ability to diminish leukotriene synthesis by zileuton, an inhibitor of 5-lipoxygenase, <i>Agents Actions Suppl.</i> , 1991, pp. 103-116, Vol. 35. (abstract only)		
	28	CARTER, et al., 5-Lipoxygenase inhibitory activity of Zileuton, <i>J. Pharmacol. & Exp. Ther</i> , 1991, pp. 929-937, Vol. 256, Vol. 3.		
	29	SUBBARAMAIAH et al., Resveratrol Inhibits the Expression of Cyclooxygenase-2 in Human Mammary and Oral Epithelial Cells, <i>Pharmaceutical Biology</i> , 1998, pp. 35-43, Vol. 36.		
	30	BINGOL et al., A Review of Terrestrial Plants and Marine Organisms Having Antiinflammatory Activity. <i>International Journal of Pharmacognosy</i> , 1995, pp. 81-97, Vol. 33, No. 2.		
	31	MARNETT et al., Arachidonic Acid Oxygenation by COX-1 and COX-2, <i>The Journal of Biological Chemistry</i> , 1999, pp. 22903-22906, Vol. 274, No. 33.		
	32	KALGUTKAR et al., Aspirin-like Molecules that Covalently Inactivate Cyclooxygenase-2, <i>Science</i> , 1998, pp. 1268-1270, Vol. 280.		
	33	KALGUTKAR, et al., Biochemically based design of cyclooxygenase-2 (COX-2) inhibitors: Facile conversion of nonsteroidal antiinflammatory drugs to potent and highly selective Cox-2 Inhibitors, <i>PNAS</i> , 2000, pp. 925-930, Vol. 97, No. 2.		

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	34	BATRISTINI et al., COX-1 and COX-2: Toward the development of More selective NSAIDs, <i>DN&P</i> , 1994, pp. 50-512, Vol. 7, No. 8.	
	35	MITCHELL et al., Cyclooxygenase-2: Regulation and Relevance in Inflammation, <i>Biochemical Pharmacology</i> , 1995, pp. 1535-1542, Vol. 50, No. 10.	
	36	WHITEHOUSE et al., Over the counter (OTC) oral remedies for arthritis and rheumatism: how effective are they?, <i>Inflammopharmacology</i> , 1999, pp. 89-105, Vol. 7, No. 2.	
	37	DEWITT et al., The Differential Susceptibility of Prostaglandin Endoperoxide H Synthases-1 and -2 to Nonsteroidal Anti-Inflammatory Drugs: Aspirin Derivatives as Selective Inhibitors, <i>Med Chem Res</i> , 1995, pp. 325-343, Vol. 5.	
	38	DUKE, J., Clippings from my COX Box, <i>Journal of Medicinal Food</i> , 1998/1999, pp. 293-298, Vol. 1, No. 4.	
	39	MCADAM et al., Systemic biosynthesis of prostacyclin by cyclooxygenase (COX)-2: The human pharmacology of a selective inhibitor of Cox-2, <i>Proc. Nat. Acad. Sci. USA</i> , 1999, pp. 272-277, Vol. 96.	
	40	O'NEILL et al., Overexpression of Human Prostaglandin G/H Synthase-1 and -2 by Recombinant Vaccinia Virus: Inhibition by Nonsteroidal Anti-Inflammatory Drugs and Biosynthesis of 15-Hydroxyeicosatetraenoic Acid, <i>Mole. Pharmacol</i> , 1994, pp. 245-254, Vol. 45.	
	41	AUERBACH et al., A Spectrophotometric Microtiter-Based Assay for the Detection of Hydroperoxy Derivatives of Linoleic Acid, <i>Anal. Biochem</i> , 1992, pp. 375-380, Vol. 201.	
	42	NOWLIN et al., A Novel Cyclic Pentapeptide Inhibits $\alpha 4\beta 1$ and $\alpha 5\beta 1$ Integrin-mediated cell Adhesion, <i>J. Biol. Chem</i> , 1993, pp. 20352-20359, Vol. 268.	
	43	STOLTENBORG et al., A fluorescent cellular adhesion assay using insect cell produced human VCAM1, <i>J. Immunological Methods</i> , 1994, pp. 59-68, Vol. 175.	
	44	WELKER et al., Glucocorticoid-Induced Modulation of Cytokine Secretion from Normal and Leukemic Human Myelomonocytic Cells, <i>International Arch of Allergy and Immunology</i> , 1996, pp. 110-115, Vol. 109.	

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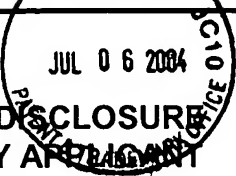
	45	KOIZUMI et al., Inhibitors of IL-2 Production and IL-2 Receptor Expression in Human Leukemic T-Cell Line, Jurkat, <i>Cellular Immunology</i> , 1986, pp. 469-475, Vol. 103.	
	46	COHEN et al., Cytokine Function, <i>Am. J. Clin. Pathol.</i> , 1996, pp. 589-598, Vol. 105., No. 5.	
	47	HENDERSON et al., Therapeutic potential of cytokine manipulation, <i>TIPS</i> , 1992, pp. 145-152, Vol. 13.	
	48	ELIAS et al., Synergistic Stimulation of Fibroblast Prostaglandin Production by Recombinant Interleukin 1 and Tumor Necrosis Factor, <i>J. Immunol.</i> , 1987, pp. 3812-3816, Vol. 138.	
	49	LENARDO et al., NF-kB: A Pleiotrophic Mediator of Inducible and Tissue-Specific Gene Control, <i>Cell</i> , 1989, pp. 227-229, Vol. 58.	
	50	MALOFF et al., Development of an RIA-based primary screen for IL-1 antagonists, <i>Clin. Chim. Acta.</i> , 1988, pp. 73-78, Vol. 180.	
	51	KARTTUMEN et al., Measurement of ligand-induced activation in single viable T cells using the <i>lacZ</i> reporter gene, <i>Proc. Nat'l. Acad. Sci.</i> , 1991, pp. 3972-3976, Vol. 88.	
	52	EMMEL et al., Cyclosporin A Specifically Inhibits Function of Nuclear Proteins Involved in T Cell Activation, <i>Science</i> , 1989, pp. 1617-1620, Vol. 246.	
	53	CHANG et al., The Influence of Chinese Traditional Medicine on the Production and Activity of Interleukin 1, <i>Chinese J. Microbiol. Immunol.</i> , 1993, pp. 15-24, Vol. 26.	
	54	MIN, K.R. et al. (-)-Epiazelechin: Cyclooxygenase-1 Inhibitor and Anti-Inflammatory Agency from Aerial Parts of <i>Celastrus orbiculatus</i> , <i>Planta Med.</i> , 1999, pp. 460-462, Vol. 65.	
	55	LEE et al. a-Viniferin: A Prostaglandin H2 Synthase Inhibitor from Root of <i>Carex humilis</i> , <i>Planta Med.</i> , 1998, pp. 195-290, Vol. 64.	
	56	NOREEN et al., Flavan-3-ols isolated from Some Medicinal Plants Inhibiting COX-1 and COX-2 Catalysed Prostaglandin Biosynthesis, 1998, pp. 520-524, Vol. 64.	
	58	DANZ et al., Identification and Isolation of the Cyclooxygenase-2 Inhibitory Principle in <i>Isatis tinctoria</i> , <i>Planta Med.</i> , 67, pp. 411-416, Vol. 67.	

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	59	TAYLOR et al., COX-1 inhibitory activity in extracts from <i>Eucomis</i> L'Herit. species, <i>J. Ethnopharmacol.</i> , 2001, pp. 257-265, Vol. 76, Nos. 2-3.	
	60	CHOU et al., The inhibitory effect of common traditional anti-rheumatic herb formulas on prostaglandin E and interleukin 2 in vitro: A comparative study with <i>Tripterygium wilfordii</i> , <i>J. Ethnopharmacol.</i> , 1998, pp. 167-171, Vol. 62, No. 2.	
	61	ZSCHOCKE et al., 5-Lipoxygenase and Cyclooxygenase Inhibitory Active Constituents from Qianghuo (<i>Notopterygium incisum</i>), <i>Planta Med.</i> , 1997, pp. 203-206, Vol. 63.	
	62	KIM et al., Effects of naturally- occurring flavonoids and biflavonoids on epidermal cyclooxygenase and lipoxygenase from guinea pigs, <i>Prostaglandins Leukot. Essent. Fatty Acids</i> , 1998 58(1); pp. 17-24.	
	63	TUNON et al., Evaluation of anti-inflammatory activity of some Swedish medicinal plants. Inhibition of prostaglandin biosynthesis and PAF-induced exocytosis, <i>J. Ethnopharmacol.</i> , 48, 1995, pp. 61-76.	
	64	YOU et al., Inhibition of Cyclooxygenase/Lipoxygenase from Human Platelets by Polyhydroxylated/Methoxylated Flavonoids Isolated from Medicinal Plants, <i>Arch. Pharm. Res.</i> , 1999, pp. 18-24, Vol. 22, No. 1.	
	65	SEGURA et al., Anti-Inflammatory Activity of Dichloromethane Extract of <i>Heterotheca inuloids</i> in Vivo and in Vitro, <i>Planta Med.</i> , 2000, pp. 553-555, Vol. 66.	
	66	DUNSTAN et al., Evaluation of some Samoan and Peruvian medicinal plants by prostaglandin biosynthesis and rat ear oedema assays, <i>J. Ethnopharmacol.</i> , 1997, pp. 35-56, Vol. 57, No. 1.	
	67	ABAD et al., The Activity of Flavonoids Extracted from <i>Tanacetum microphyllum</i> DC. (Compositae) on Soybean Lipoxygenase and Prostaglandin Synthetase, <i>Gen. Pharmacol.</i> , 1995, pp. 815-819, Vol. 26, No. 4.	
	68	ABAD et al., Anti-inflammatory Activity of Hydroxyachillin, a Sesquiterpene Lactone from <i>Tanacetum microphyllum</i> , <i>Planta Med.</i> , 1994, pp. 228-231, Vol. 60.	
	69	WANG et al., Antioxidant and Antiinflammatory Activities of Anthocyanins and Their Aglycon, Cyanidin, from Tart Cherries, <i>J. Nat. Prod.</i> , 1999, pp. 294- 296, Vol. 62, No. 2.	

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70	RESCH et al., Further Phenols and Polyacetylenes from the Rhizomes of <i>Atractylodes lancea</i> and their Anti-Inflammatory Activity, <i>Planta Med.</i> , 2001, pp. 437-442. Vol. 67.	
71	JAGER et al., Screening of Zulu medicinal plants for prostaglandin synthesis inhibitors, <i>J. Ethnopharmacol.</i> , 1996, pp. 95-100, Vol. 52, No. 2.	
72	RINGBOM et al., COX-2 Inhibitory Effects of Naturally Occurring and Modified Fatty Acids, <i>J. Nat. Prod.</i> , 2001, pp., 745-749, Vol. 64, No. 6.	
73	NOREEN et al., Development of a Radiochemical Cyclooxygenase-1 and -2 in Vitro Assay for Identification of Natural Products as Inhibitors of Prostaglandin Biosynthesis, <i>J. Nat. Prod.</i> , 1998, Vol. 61, No. 1.	
74	MOON et al., A new class of COX-2 inhibitor, rutaecarpine from <i>Evodia rutaecarpa</i> , <i>Inflamm. Res.</i> , 1999, Vol. 48, pp. 621-625.	
75	REDDY et al., Selective Inhibition of Cyclooxygenase-2 by C-Phycocyanin, a Biliprotein from <i>Spirulina platensis</i> , <i>Biochem. Biophys. Res. Commun.</i> , 2000, Vol. 277, No. 3.	
76	LIU et al., Inhibitory Effects of <i>Angelica pubescens f. biserrata</i> on 5-Lipoxygenase and Cyclooxygenase, <i>Planta Med.</i> , 1998, pp. 525-529, Vo. 64.	
77	NEWMARK et al. (2000) Beyond Aspirin: Nature's Answer to Arthritis, Cancer, and Alzheimer's Disease. Hohm Press, Prescott, AZ. (Table of Contents only).	
78	LAVALLE, James B. (2001) The COX-2 Connection: Natural Breakthrough Treatments for Arthritis, Alzheimer's, and Cancer. Healing Arts Press. (Table of Contents only).	
79	TYLER, Varro E., Phytomedicines: Back to the Future, <i>J. Nat. Prod.</i> , 1999, pp. 1589-1592.	
80	MICHALUART et al, Inhibitory Effects of Caffeic Acid Phenethyl Ester on the Activity and Expression of Cyclooxygenase-2 in Human Oral Epithelial Cells and in a Rat Model of Inflammation, <i>Cancer Res.</i> , 1999, Vol. 59, pp. 2347-2352.	
81	SUBBARAMAIAH et al., Ursolic acid inhibits cyclooxygenase-2 transcription in human mammary epithelial cells, <i>Cancer Res.</i> , 2000, Vol. 60, pp. 2399-2404.	
82	SUBBARAMAIAH et al, Resveratrol inhibits cyclooxygenase-2 transcription in human mammary epithelial cells, <i>Ann. N.Y. Acad. Sci.</i> , 1999, Vol. 889, pp. 214-223.	

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PTO/SB/08A INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)				Complete if Known	
				Application Number	10/022,320
				Filing Date	12-13-2001
				Confirmation Number	4956
				First Named Inventor	M. Obukowicz
				Group Art Unit	1614
Sheet	8	of	9	Attorney Docket No.	3412/1A/US (PHA 4140.6)

	83	SUBBARAMAIIH et al., Resveratrol inhibits cyclooxygenase-2 transcription and activity in phorbol ester-treated human mammary epithelial cells, <i>J. Biol. Chem.</i> , 1998, 273(34):21875-21882.	
	84	WILLIAMS et al. The flavonoids of <i>Tanacetum parthenium</i> and <i>T. vulgare</i> and their anti-inflammatory properties, <i>Phytochemistry</i> , 1999, pp. 417-423, Vol. 51, No. 3.	
	85	REDDY et al., Studies on Anti-Inflammatory Activity of Spice Principles and Dietary n-3 Polyunsaturated Fatty Acids on Carrageenan-Induced Inflammation in Rats, <i>Ann. Nutr. Metab.</i> , 1994, pp. 349-358.	
	86	JUERGENS, U.R., et. al., The Anti-Inflammatory Activity of L-Menthol Compared to Mint Oil in Human Monocytes In Vitro: A Novel Perspective For Its Therapeutic Use in Inflammatory Diseases, <i>Eur. J. Med. Res.</i> , 1998, pp. 539-545.	
	87	JUERGENS, U.R., et. al., Inhibition of Cytokine Production And Arachidonic Acid Metabolism By Eucalyptol (1.8-Cineole) In Human Blood Monocytes In Vitro, <i>Eur. J. Med. Res.</i> , 1998, pp. 508-510.	
	88	HAQQI et al., Prevention of collagen-induced arthritis in mice by a polyphenolic fraction from green tea, <i>Proc. Natl. Acad. Sci. USA</i> , 1999, pp. 4524-4529, Vol. 96.	
	89	ALANKO et al., Modulation of arachidonic acid metabolism by phenols: Relation to their structure and anti-oxidant/pro-oxidant properties, <i>Free Radical Biology and Medicine</i> , 1999, pp. 193-201, Vol. 26.	
	90	ZHANG et. al., Curcumin inhibits cyclooxygenase-2 transcription in bile acid- and phorbol ester-treated human gastrointestinal epithelial cells, <i>Carcinogenesis</i> , 1999, Vol. 20, No. 3.	
	91	LINDSEY et al., Screening of plants used by Southern African traditional healers in the treatment of dysmenorrhoea for prostaglandin-synthesis inhibitors and uterine relaxing activity, <i>Journal of Ethnopharmacology</i> , 1999, pp. 9-14, Vol. 64.	
	92	JANG et al., Cancer chemopreventive activity of resveratrol, a natural product derived from grapes, <i>Science</i> , 1997, pp. 218-220, Vol. 275.	
	93	GIERSE et al., A single Amino Acid Difference Between Cyclooxygenase-1 (COX-1) and -2 (COX-2) Reverses the Selectivity of COX-2 Specific Inhibitors, <i>The Journal of Biological Chemistry</i> , 1996, pp. 15810-15814, Vol. 271, No. 26.	

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	94	Nexrutine brochure for Remedief, Triple Action Pain Management with Natural Cox-2 Inhibitor. Undated	
	95	Hawley's Condensed Chemical Dictionary, 13th Edition, Revised by Lewis, 1997, page 736.	
	96	J. Heterocyclic Chem. 1982, pp. 193-200, Vol. 19, No. 1.	

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